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2	484	((plasminogen or plasmin) near2 (fragment))	USPAT; US-PGPUB; EPO; JPO; DERWENT	2004/10/29 15:42
3	296	((plasminogen or plasmin) near2 (fragment)) NOT ((plasminogen or plasmin) near3 (inhibitor))	USPAT; US-PGPUB; EPO; JPO; DERWENT	2004/10/29 15:42
4	181	((plasminogen or plasmin) near2 (fragment)) NOT ((plasminogen or plasmin) near3 (inhibitor)) ) NOT ((plasminogen or plasmin) near3 (activator))	USPAT; US-PGPUB; EPO; JPO; DERWENT	2004/10/29 15:42
5	180	((plasminogen or plasmin) adj (fragment))	USPAT; US-PGPUB; EPO; JPO; DERWENT	2004/10/29 15:42
6	77	((plasminogen or plasmin) adj (fragment)) NOT ((plasminogen or plasmin) near3 (inhibitor))	USPAT; US-PGPUB; EPO; JPO; DERWENT	2004/10/29 15:42
7	49	((plasminogen or plasmin) adj (fragment)) NOT ((plasminogen or plasmin) near3 (inhibitor)) ) NOT ((plasminogen or plasmin) near3 (activator))	USPAT; US-PGPUB; EPO; JPO; DERWENT	2004/10/29 15:42

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=> s plasmin (4A) (peptide antibody)

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L2 ANSWER 1 OF 4 CAPLUS COPYRIGHT 2004 ACS on STN DUPLICATE 1

AN 1997:181681 CAPLUS

DN 126:223529

TI Preparation and proteolytic degradation of a macromolecular protein  
 complex from fibrinogen  
 IN Lipinski, Boguslaw  
 PA Lipinski, Boguslaw, USA  
 SO U.S., 7 pp.  
 CODEN: USXXAM  
 DT Patent  
 LA English  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 5607841	A	19970304	US 1994-262607	19940620
PRAI	US 1994-262607		19940620		

AB A method for preparing a macromol. protein complex (MPC) from fibrinogen in  
 human plasma by the steps of (NH<sub>4</sub>)<sub>2</sub>SO<sub>4</sub> precipitation, dialysis, and gel  
 filtration

is disclosed. The isolated MPC may be degraded by contacting with  
 trypsin. The isolated MPC inhibited fibrinolysis induced with plasminogen  
 but not with plasmin. Elimination of the MPC by means of chondroitin  
 sulfate A restored normal fibrinolysis. An antibody to fibrin-binding  
 peptides which are produced by trypsin degradation of MPC was conjugated to  
 plasmin. The anti-MPC **peptide antibody/**  
**plasmin** conjugate has the capacity to bind to MPC-rich thrombus  
 and degrade it without activation of fibrin-bound plasminogen.

L2 ANSWER 2 OF 4 BIOSIS COPYRIGHT (c) 2004 The Thomson Corporation. on STN  
 DUPLICATE 2

AN 1997:390155 BIOSIS

DN PREV199799689358

TI The cluster of basic amino acids in vitronectin contributes to its binding  
 of plasminogen activator inhibitor-1: Evidence from thrombin-, elastase-  
 and **plasmin**-cleaved vitronectins and anti-**peptide**  
**antibodies.**

AU Gechtman, Zeev; Belleli, Adina; Lechpammer, Stanislav; Shaltiel, Shmuel  
 [Reprint author]

CS Dep. Biological Regulation, Weizmann Inst. Science, IL-76100 Rehovot,  
 Israel

SO Biochemical Journal, (1997) Vol. 325, No. 2, pp. 339-349.  
 ISSN: 0264-6021.

DT Article

LA English

ED Entered STN: 10 Sep 1997

Last Updated on STN: 10 Sep 1997

AB Derivatives of vitronectin obtained by specific cleavage at its cluster of  
 basic amino acids with thrombin, elastase and plasmin are shown to have a  
 decreased ability to bind plasminogen activator inhibitor-1 (PAI-1). The  
 identification and localization of the segment involved in the binding of  
 PAI-1 (Lys-348-Arg-379) were carried out by purification of these cleaved  
 vitronectins and their subsequent structural characterization (sequence  
 analysis, phosphorylation of Ser-378 with cAMP-dependent protein kinase  
 and immunostaining with peptide-specific antibodies), then measurement of  
 the vitronectin-PAI-1 interaction by (a) a two-phase system (ELISA); (b)  
 co-precipitation of the vitronectin-PAI-1 complex out of solution, and (c)  
 analysis of the stereospecific interaction between the active conformation  
 of PAI-1 and a peptide derived from the above-mentioned cluster; this  
 interaction occurs when the peptide is composed of all-L-amino acids but  
 not when it is composed of all-D-amino acids. Our results explain why  
 workers who have used immobilized vitronectin to study this interaction  
 could not have observed the involvement of the cluster of basic amino  
 acids in PAI-1 binding, since the immobilization of vitronectin is shown  
 to render this cluster inaccessible for interaction. We propose that  
 vitronectin binds active PAI-1 by interaction via amino acid residues that  
 originate from distal locations in the N- and C-termini of vitronectin.



L2 ANSWER 3 OF 4 WPINDEX COPYRIGHT 2004 THE THOMSON CORP on STN  
 AN 1994-209452 [26] WPINDEX  
 DNN N1994-164933 DNC C1994-095856  
 TI Synthetic peptide(s) and antibodies against fragment E - derived from plasmin-cleaved fibrinogen useful for therapy of disturbances in the fibrinolytic system.  
 DC B04 D16 S03  
 IN KRAUS, M; STUBER, W; STUEBER, W  
 PA (BEHW) BEHRINGWERKE AG; (DADE-N) DADE BEHRING MARBURG GMBH  
 CYC 18  
 PI DE 4242736 A1 19940623 (199426)\* 33  
 EP 605797 A1 19940713 (199427) GE 45  
 R: AT BE CH DE DK ES FR GB IT LI LU NL PT SE  
 AU 9352435 A 19940630 (199430)  
 CA 2111645 A 19940618 (199432)  
 JP 06256388 A 19940913 (199441) 19  
 US 5599678 A 19970204 (199711) 30  
 AU 676859 B 19970327 (199721)  
 EP 605797 B1 19990317 (199915) GE  
 R: AT BE CH DE DK ES FR GB IT LI LU NL PT SE  
 DE 59309458 G 19990422 (199922)  
 ES 2129487 T3 19990616 (199930)  
 US 5981697 A 19991109 (199954)  
 US 6441141 B1 20020827 (200259)  
 ADT DE 4242736 A1 DE 1992-4242736 19921217; EP 605797 A1 EP 1993-119574 19931209; AU 9352435 A AU 1993-52435 19931215; CA 2111645 A CA 1993-2111645 19931216; JP 06256388 A JP 1993-344306 19931217; US 5599678 A US 1993-166930 19931215; AU 676859 B AU 1993-52435 19931215; EP 605797 B1 EP 1993-119574 19931209; DE 59309458 G DE 1993-509458 19931209, EP 1993-119574 19931209; ES 2129487 T3 EP 1993-119574 19931209; US 5981697 A Div ex US 1993-166930 19931215, US 1996-727045 19961008; US 6441141 B1 Div ex US 1993-166930 19931215, Div ex US 1996-727045 19961008, US 1999-408172 19990929  
 FDT AU 676859 B Previous Publ. AU 9352435; DE 59309458 G Based on EP 605797; ES 2129487 T3 Based on EP 605797; US 5981697 A Div ex US 5599678; US 6441141 B1 Div ex US 5599678, Div ex US 5981697  
 PRAI DE 1992-4242736 19921217  
 AB DE 4242736 A UPAB: 19940817  
 Claimed are (a) synthetic peptides of antigenic aminoacid sequences which at least partially correspond to the carboxyl terminal ends of fragment E obtd. after plasmin-cleavage of fibrinogen, (b) synthetic peptides containing at least one of (i) Leu-Phe-Glu-Tyr-Gln-Lys-OH, (ii) Tyr-Met-Tyr-Leu-Leu-Lys-OH (iii) Val-Lys-Glu-Leu-Ile-Lys-OH, and (iv) His-Gln-Val-Glu-Asn-Lys-OH (c) antibodies (ABs) reacting with (a).  
 USE/ADVANTAGE - (a) or (c) are used for the immunochemical detection of oligomer peptides of fibrinogen or fibrin cleavage prods. having at least one immunochemically identical epitope either in a heterogenous (enzyme immuno assay) or homogeneous (nephelometric or turbidimetric test) immuno assay. The ABs and peptides are used in the treatment of fibrinolytic system disturbances. The invention provides antigens from which specific AB for fibrinogen-or fibrin cleavage prods. can easily be produced for an exact quantitative measurement of the fibrinolytic activity in biological fluids. In contrast to known method cross-reactivity is avoided since the ABs only react with fragment E but do not recognise native fibrinogen or fibrin.  
 Dwg.1/16

L2 ANSWER 4 OF 4 BIOSIS COPYRIGHT (c) 2004 The Thomson Corporation. on STN  
 DUPLICATE 3  
 AN 1992:139004 BIOSIS  
 DN PREV199293073229; BA93:73229  
 TI ANTI-IDIOTYPIC ANTIBODIES AGAINST AN ANTIBODY TO THE PLATELET GLYCOPROTEIN

GP IIB-IIIA COMPLEX MIMIC GP IIB-IIIA BY RECOGNIZING FIBRINOGEN.

AU ABRAMS C S [Reprint author]; RUGGERI Z M; TAUB R; HOXIE J A; NAGASWAMI C;  
WEISEL J W; SHATTIL S J

CS HEMATOL ONCOL SECTION, HOSPITAL UNIVERSITY PENNSYLVANIA, 3400 SPRUCE ST,  
PHILADELPHIA, PA 19104, USA

SO Journal of Biological Chemistry, (1992) Vol. 267, No. 4, pp. 2775-2785.  
CODEN: JBCHA3. ISSN: 0021-9258.

DT Article

FS BA

LA ENGLISH

ED Entered STN: 12 Mar 1992  
Last Updated on STN: 12 Mar 1992

AB Binding of the adhesive ligand fibrinogen and the monoclonal antibody PAC1 to platelet glycoprotein (GP) IIb-IIIa is dependent on cell activation and inhibited by Arg-Gly-Asp (RGD)-containing peptides. Previously, we identified a sequence in a hypervariable region of PAC1 ( $\mu$ -CDR3) that mimics the activity of the antibody. Here we examine whether monoclonal antibodies to this idiotypic determinant in PAC1 can mimic GP IIb-IIIa by binding to fibrinogen. Mice were immunized with a peptide derived from the  $\mu$ -CDR3 of PAC1. Four antibodies were obtained that recognized fibrinogen as well as a recombinant form of the variable region of PAC1. However, they did not bind to other RGD-containing proteins, including von Willebrand factor, fibronectin, and vitronectin. Several studies suggested that these anti-PAC1 peptide antibodies were specific for GP IIb-IIIa recognition sites in fibrinogen. Three such sites have been proposed: two RGD-containing regions in the A $\alpha$  chain, and the COOH terminus of the  $\gamma$  chain ( $\gamma$ 400-411). Two of the antibodies inhibited fibrinogen binding to activated platelets, and all four antibodies bound to the fibrinogen A $\alpha$  chain on immunoblots. Antibody binding to immobilized fibrinogen was partially inhibited by monoclonal antibodies specific for the two A $\alpha$  chain RGD regions. However, the anti-PAC1 **peptide antibodies** also bound to **plasmin**-derived fibrinogen fragments X and D100, which contain  $\gamma$ 400-411 but lack one or both A $\alpha$ RGD regions. This binding was inhibited by an antibody specific for  $\gamma$ 400-411. When fragment D100 was converted to D80, which lacks  $\gamma$ 400-411, antibody binding was reduced significantly ( $p < 0.01$ ). Electron microscopy of fibrinogen-antibody complexes confirmed that each antibody could bind to sites on the A $\alpha$  and  $\gamma$  chains. These studies demonstrate that certain anti-PAC1 peptide antibodies mimic GP IIb-IIIa by binding to platelet recognition sites in fibrinogen. Furthermore, they suggest that the  $\gamma$ 400-411 region of fibrinogen may exist in a conformation similar to that of an A $\alpha$ RGD region of the molecule.